

Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

### **REMARKS**

Applicants respectfully request reconsideration of the application, as amended, in view of the following remarks.

Notably, Applicants have added **new Claims 31 and 32** defining the additive as ethylene glycol.

The present invention as set forth in **Claim 1** relates to a **porous hollow fiber membrane** obtained by a dry-wet spinning method or a wet spinning method from a spinning dope while using the following components:

a spinning dope containing a base polymer as a material for forming said porous hollow fiber membrane,

**an additive for facilitating a phase separation of said spinning dope,**

a solvent compatible with both, said base polymer and said additive, and

a mass of microparticles insoluble in said solvent, wherein said microparticles are uniformly dispersed in a liquid medium and have an average particle size within the range of 1 to 20  $\mu\text{m}$ , and

a coagulating liquid for forming the hollow fiber membrane,

to obtain a spun hollow fiber membrane; and

**extracting and removing said additive** and said microparticles by immersing said spun hollow fiber membrane into an extracting solution effective to dissolve said microparticles, but ineffective to dissolve said base polymer;

wherein said hollow fiber membrane has a permselectivity; wherein **a particle cutoff is within the range of 1 to 10  $\mu\text{m}$ ; and wherein a pure water permeate flow is equal to or**

**higher than 30,000 L/m<sup>2</sup>/hr/100kPa.**

**Claim 6** relates to a method of making a porous hollow fiber membrane.

For the Examiner's convenience, Applicants summarize the main arguments which Applicants have previously made.

1) Parham et al, Ruckenstein et al (US 5,993,661) and Stengaard, neither disclose nor suggest the claimed membrane having a **particle cutoff of 1 to 10 $\mu$ m** or the claimed **water permeate flow** or that such can be attained using an **additive**.

2) Since the proposed modification of Parham et al (increase of particle cutoff) would render the prior art invention being modified unsatisfactory for its intended purpose (filtration of LDL cholesterol from blood), there is no suggestion or motivation to make the proposed modification (*In re Gordon*, 733 F.2d 900, 221 USPQ 1125 (Fed. Cir. 1984)).

3) Even if Parham et al were combined with Ruckenstein et al and/or Stengaard, the present invention would not result because Ruckenstein et al disclose the use of much larger silica particles, for example, 15 to 40 microns.

4) Further, the use of **an additive** such as ethylene glycol mentioned in the specification on page 29, line 1 (Example 1) facilitates a phase separation of the spinning dope so that the pore size resulting in the **particle cutoff and water permeate flow** claimed in Claims 1 and 6 can be attained.

5) The **Rule 132 Declaration** filed **December 1, 2003**, shows the influence of the additive on the pore size. Notably, **without** the use of the **additive**, the claimed **water permeate flow of equal to or higher than 30,000 L/m<sup>2</sup>/hr/100kPa cannot be achieved**.

Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

Applicants take notice that the Examiner has not at all addressed the Rule 132 Declaration in the Office Action of February 19, 2004. Accordingly, Applicants attach a copy of the Declaration herewith so that the Examiner can respond to the presented data.

The Examiner argues that even though Parham et al “does not teach the pore size or cast-off of the membrane intended for a particular separation process”, it “teaches the skilled in the art how to make membranes from the same process with pore sizes larger than the desired pore size...” In particular, it is the Examiner’s opinion that it would have been obvious to increase the particle size to obtain a greater pore size. (Office Action of May 30, 2003, page 3, lines 4-9). Specifically, the Examiner now cites JPA 58-91822 because it is her opinion that it “clearly teaches the effect of silica size particles in the final membrane pore size and the increase in pore size of the formed membrane by the increase in silica particle size.” (Office Action of February 19, 2004, page 2, lines 5-7 from the bottom).

However, JPA 58-91822 discloses at page 7, left lower column, lines 18-20 that, when the particle size is greater than 5 microns, the membrane tends to be **non-uniform** in pore size with **large voids** embedded. Thus, membrane forming becomes difficult or impossible when the particle size reaches over 5 microns. Further, Example 2 in JPA 58-91822 describes the use of finely-powdered silica with an average particle size of **3.5 microns** and the resultant hollow membranes have a rejection of particles of 2000 Å. The value 2000 Å corresponds to particle cutoff of 0.2 microns. Thus, it can be reasonably estimated that the use of **particle size** of the upper limit, **5 microns**, of JPA 58-91822 results in hollow membranes having a rejection of particle of around 3000 Å or **0.3 microns** which is still **smaller than the range of 1 to 10 microns claimed in Claims 1 and 6** of the present

Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

invention. Since the use of particles having greater particle size than 5 microns renders membrane forming difficult or impossible according to JPA 58-91822, it can be concluded that the pore size or **particle cut-off range of greater than 0.3 microns can not be attained in the prior art**. Accordingly, JPA 58-91822 does not disclose nor suggest the present invention at all.

Further, the Examiner argues that Parham et al disclose the addition of an additive and that “the effect in facilitating phase separation is inherent of the additive, e.g. PEG.” (Office Action of February 19, 2004, page 4, lines 10-13).

Parham et al utilizes polyethylene glycol (PEG) or hydrophilic polymer to modify the structure and surface characteristics of the polysulfone membrane ( Parham et al, col. 5, lines 14-22). There is no disclosure or suggestion that the PEG is used to enlarge pore size. Furthermore, in Parham et al only Example 2 employs a hydrophilic polymer which is added not during the membrane forming process but **after the hollow membrane are formed**. Thus, the PEG is used to provide the membrane with hydrophilic properties by leaving the PEG in the membrane. In fact, Parham et al discloses at col. 5, lines 23 and 24 : “**The additional polymer or prepolymer becomes an integral part of the membrane structure.**” In contrast, in Claims 1 and 6 of the present invention, the additive is **extracted and removed. Thus, the resulting membranes are very different.**

In the following, Applicants provide a detailed explanation of the arguments 1)-5) which were summarized above.

Parham et al (US 5,258,149) disclose a membrane having a pore diameter of **about**

Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

**0.1 to about 0.7 microns** (Parham et al, col. 7, lines 49 and 50) which is much smaller than the claimed particle cutoff of **1 to 10 $\mu$ m**. There is no suggestion or motivation to go to a higher pore size as this would be detrimental to the filtration capability of the membrane for LDL cholesterol. **If the proposed modification would render the prior art invention being modified unsatisfactory for its intended purpose, then there is no suggestion or motivation to make the proposed modification** (*In re Gordon*, 733 F.2d 900, 221 USPQ 1125 (Fed. Cir. 1984)). Parham et al have chosen a specific pores size so that LDL cholesterol can be filtered from whole blood. Parham et al state at col. 7, lines 41-50 as follows:

“ The dimensional and porosity characteristics of the membranes of this invention are such that **LDL-C can pass through the fiber wall but most blood cells do not. Hemolysis** occurs if numerous blood cells pass through the fibers, **which is highly undesirable.....**Generally speaking, membranes can be prepared which possess a pore diameter of between about 0.1 microns to about 0.7 microns, preferably between 0.4 and 0.65 microns.”

Thus, Parham et al clearly teach away from larger pore sizes as they would lead to undesirable hemolysis. If the pore size of Parham et al were increased, the membrane would be completely useless. Since an increase in pore size of Parham et al would render the membrane completely useless, there cannot be any suggestion or motivation to increase the pore size.

In addition, since the membrane of Parham et al does not have the claimed pore size it **cannot have the claimed water permeate flow**. This is further shown by Comparison Example 2 at page 33 of the specification. Here a membrane with a particle cutoff of **0.85**

Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

**micron** was prepared. The corresponding pure water permeate flow is only **22,000 L/m<sup>2</sup>/hr/100kPa**, which is much lower than the **claimed** pure water permeate flow of **30,000 L/m<sup>2</sup>/hr/100kPa**. A smaller pore size such as disclosed in Parham et al would yield an even smaller pure water permeate flow.

Furthermore, one feature of the present invention is to use **an additive** such as ethylene glycol mentioned in the specification on page 29, line 1 (Example 1). The pore size resulting in the particle cutoff claimed in Claims 1 and 6 can be attained by using the claimed additive for facilitating a phase separation of the spinning dope. As mentioned in the specification at page 11, lines 22-24, addition of one or more additives is effective to enhance a phase separation of the spinning dope, which eventually results information of the hollow fiber membrane having a relatively large pore size. Furthermore, as mentioned on page 12, lines 1 to 7, an appropriate selection of the additive amount and mixture of the microparticles contributes further in providing large size pores. Moreover, Applicants again wish to draw the Examiner's attention to the **Rule 132 Declaration filed December 1, 2003**, showing the influence of the additive on the pore size.

The results obtained **with additive (Examples 1, 3 and 5)** or **without additives (Examples 2, 4 and 6)** are summarized in the Tables below:

Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

|                             |   |              |                                       |        |
|-----------------------------|---|--------------|---------------------------------------|--------|
|                             | <b>Example 1</b><br>polysulfone         | 20 wt%       | <b>Example 2</b><br>polysulfone       | 20 wt% |
| spinning dope               | <b>ethylene glycol</b>                  | <b>6 wt%</b> | <b>ethylene glycol</b>                | --     |
|                             | silicone oxide (4.5 $\mu$ m)            | 18 wt%       | silicone oxide (4.5 $\mu$ m)          | 18 wt% |
|                             | N,N-dimethyl acetoamide                 | 54 wt%       | N,N-dimethyl acetoamide               | 60 wt% |
| pure water<br>permeate flow | <b>135000 L/m<sup>2</sup>/hr/100KPa</b> |              | <b>985 L/m<sup>2</sup>/hr/100KPa</b>  |        |
| particle cutoff             | <b>2.4 <math>\mu</math>m</b>            |              | <b>0.9 <math>\mu</math>m</b>          |        |
|                             | <b>Example 3</b><br>polysulfone         | 20 wt%       | <b>Example 4</b><br>polysulfone       | 20 wt% |
| spinning dope               | <b>ethylene glycol</b>                  | <b>6 wt%</b> | <b>ethylene glycol</b>                | --     |
|                             | silicone oxide (11 $\mu$ m)             | 20 wt%       | silicone oxide (11 $\mu$ m)           | 20 wt% |
|                             | silicone oxide (4.5 $\mu$ m)            | 2 wt%        | silicone oxide (4.5 $\mu$ m)          | 2 wt%  |
|                             | N,N-dimethyl acetoamide                 | 52 wt%       | N,N-dimethyl acetoamide               | 58 wt% |
| pure water<br>permeate flow | <b>520000 L/m<sup>2</sup>/hr/100KPa</b> |              | <b>1746 L/m<sup>2</sup>/hr/100KPa</b> |        |
| particle cutoff             | <b>5.0 <math>\mu</math>m</b>            |              | <b>2.1 <math>\mu</math>m</b>          |        |
|                             | <b>Example 5</b><br>polysulfone         | 20 wt%       | <b>Example 6</b><br>polysulfone       | 20 wt% |
| spinning dope               | <b>ethylene glycol</b>                  | <b>4 wt%</b> | <b>ethylene glycol</b>                | --     |
|                             | silicone oxide (1.5 $\mu$ m)            | 14 wt%       | silicone oxide (1.5 $\mu$ m)          | 14 wt% |
|                             | N,N-dimethyl acetoamide                 | 62 wt%       | N,N-dimethyl acetoamide               | 66 wt% |
| pure water<br>permeate flow | <b>39000 L/m<sup>2</sup>/hr/100KPa</b>  |              | <b>5567 L/m<sup>2</sup>/hr/100KPa</b> |        |
| particle cutoff             | <b>1.2 <math>\mu</math>m</b>            |              | <b>0.9 <math>\mu</math>m</b>          |        |

From the results shown in the above Tables, it is clear that the additive in the present invention contributes greatly in obtaining the claimed pure water permeate flow and the claimed particle cutoff. Thus, superior results are obtained using the additive according to

Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

the present invention. Applicants wish to draw the Examiner's special attention to the **electron microphotographs attached to the Rule 132 Declaration** which further illustrate the superior properties of the membranes according to the present invention.

In order to cure the defects of Parham et al, the Examiner cites Ruckenstein et al (US 5,993,661). However, Ruckenstein et al do not cure these defects because they also fail to disclose or suggest the claimed membrane having a particle cutoff of 1 to 10 $\mu$ m or the claimed water permeate flow. Ruckenstein et al disclose the use of silica particles of large size, for example, 15 to 40 microns. This particle size is much larger than the claimed particle size. In addition, if the pore size of Parham et al (0.1 microns to about 0.7microns) was modified to a pore size of 15-40 microns using the particle size of Ruckenstein et al (Ruckenstein et al, col. 4, line 48), **Parham et al's membrane would be rendered useless as it would pass not only the LDL-C but also most blood cells.** Thus, there is no suggestion or motivation to modify the pore size of Parham et al to that of Ruckenstein et al.

In addition, Ruckenstein et al fail to disclose or suggest the superior properties obtained with the claimed additive.

Even further, Ruckenstein et al uses a **flat membrane** and not a porous hollow membrane as claimed. Membrane forming conditions for the hollow membrane are severer than those for the flat membrane of Ruckenstein et al and accordingly, in the hollow membrane pores formed by the particles tend to shrink resulting in a much smaller size of the pores.

Stengaard also does not cure the defects of Parham et al and Ruckenstein et al because it also fails to disclose or suggest the claimed membrane having a particle cutoff of 1 to 10 $\mu$ m



Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

or the claimed water permeate flow. As discussed above, there is no motivation to modify the pore size of Parham et al. In addition, Stenggaard fails to disclose or suggest the superior properties obtained with the claimed additive.

Therefore, the rejection of Claims 1-4, 6-8, and 29-30 under 35 U.S.C. §103(a) as being unpatentable over Parham et al (US 5,258,149) or Parham et al (US 5,258,149) in view of Ruckenstein et al (US 5,993,661) and the rejection of Claims 5 and 9 under 35 U.S.C. §103(a) as being unpatentable over Parham et al (US 5,258,149) or Parham et al (US 5,258,149) in view of Ruckenstein et al (US 5,993,661), and further in view of Stengard (US 5,019,261) are believed to be unsustainable as the present invention is neither anticipated nor obvious and withdrawal of these rejections is respectfully requested.

Applicants also appreciate the indication of a rejoinder of Claims 10-28 after allowability of the above Claims has been determined (see Office Action of November 22, 2002).

Application No.: 09/884,084  
Amendment Dated: May 19, 2004  
Reply to Office Action of: February 19, 2004

This application presents allowable subject matter, and the Examiner is kindly requested to pass it to issue. Should the Examiner have any questions regarding the claims or otherwise wish to discuss this case, he is kindly invited to contact Applicants' below-signed representative, who would be happy to provide any assistance deemed necessary in speeding this application to allowance.

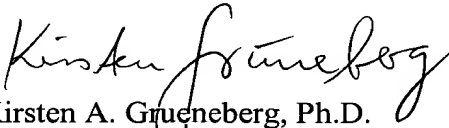
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